L Number	Hits	Search Text	DB	Time stamp
1	0	protease near4 (anhydridized or anhydridize or anhydridization)	USPAT; US-PGPUB; EPO; JPO; DERWENT	2003/07/07 12:11
7	0	enzyme near4 (anhydridized or anhydridize or anhydridization)	USPAT; US-PGPUB; EPO; JPO; DERWENT	2003/07/07 12:12
13	0	activity near4 (anhydridized or anhydridize or anhydridization)	USPAT; US-PGPUB; EPO; JPO; DERWENT	2003/07/07 12:12
19	51	anhydridized or anhydridize or anhydridization	USPAT; US-PGPUB; EPO; JPO; DERWENT	2003/07/07 12:12
25	34	(anhydridized or anhydridize or anhydridization) and (protease or enzyme or activity)	USPAT; US-PGPUB; EPO; JPO; DERWENT	2003/07/07 12:13
31	4	(anhydridized or anhydridize or anhydridization) and protease	USPAT; US-PGPUB; EPO; JPO; DERWENT	2003/07/07 12:17
37	4	((anhydridized or anhydridize or anhydridization) and protease) and (anhydridized or anhydridize or anhydridization)	USPAT; US-PGPUB; EPO; JPO; DERWENT	2003/07/07 12:17
43	11	(anhydridized or anhydridize or anhydridization) and enzyme	USPAT; US-PGPUB; EPO; JPO; DERWENT	2003/07/07 12:17

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NEWS 43   Jun 06   PASCAL enhanced with additional data
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TI      Aza- and polyaza-naphthalenyl carboxamides useful as HIV integrase
        inhibitors
IN      Anthony, Neville J., Hatfield, PA, UNITED STATES
        Gomez, Robert P., Perkasie, PA, UNITED STATES
        Young, Steven D., Lansdale, PA, UNITED STATES
        Egbertson, Melissa, Ambler, PA, UNITED STATES
        Wai, John S., Harleysville, PA, UNITED STATES
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        Perlow, Debra S., East Greenville, PA, UNITED STATES
        Bennett, Jennifer J., East Norriton, PA, UNITED STATES
        Funk, Timothy W., Ephrata, PA, UNITED STATES
PI      US 2003055071      A1   20030320
AI      US 2001-973853     A1   20011010 (9)
PRAI    US 2000-239707P     20001012 (60)
        US 2001-281656P     20010405 (60)
DT      Utility
FS      APPLICATION
LREP    MERCK AND CO INC, P O BOX 2000, RAHWAY, NJ, 070650907
CLMN    Number of Claims: 36
ECL     Exemplary Claim: 1
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AN 1999:142175 BIOSIS
DN PREV199900142175
TI Symmetrical **anhydride**-type serine **protease** inhibitors:
Structure-activity relationship studies of human chymase inhibitors.
AU Iijima, Kiyoko; Katada, Jun; Hayashi, Yoshio (1)
CS (1) Life Sci. Res. Center, Advanced Technol. Res. Lab., Nippon Steel Corp., 3-35-1 Ida, Nakahara-ku, Kawasaki 211-0035 Japan
SO Bioorganic & Medicinal Chemistry Letters, (Feb. 8, 1999) Vol. 9, No. 3, pp. 413-418.
ISSN: 0960-894X.
DT Article
LA English
AB We prepared a potent and relatively selective human chymase inhibitor 9 (-), based on the study of SAR of a symmetrical **anhydride**-type serine **protease** inhibitor 1. Kinetic studies suggested that 9 (-) reacts with the Ser residue at the active site of the enzyme, forming a stable acyl enzyme complex. We also showed the importance of the tri-substituted beta-amino acid structure for the potent anti-enzymatic activity.

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AN 1999:153482 BIOSIS
DN PREV199900153482
TI Detection of an anhydride intermediate in the carboxypeptidase A catalyzed hydrolysis of a peptide substrate by solid state NMR spectroscopy and its mechanistic implication.
AU Lee, Hee Cheon (1); Ko, Young Ho; Baek, Seung Bin; Kim, Dong H. (1)
CS (1) Dep. Chem. and Center Biofunctional Molecules, Pohang Univ. Sci. and Technol., San 31 Hyojadong, Pohang 790-784 South Korea
SO Bioorganic & Medicinal Chemistry Letters, (Dec. 1, 1998) Vol. 8, No. 23, pp. 3379-3384.
ISSN: 0960-894X.
DT Article
LA English
AB We have detected an **anhydride** intermediate in the CPA catalyzed **proteolytic** reaction of Gly-Tyr. It appears that since the zinc-bound water molecule which is believed to attack the scissile amide carbonyl carbon in the hydrolysis reaction is excluded by the N-terminal amino group of Gly-Tyr, the carboxylate of Glu-270 becomes to attack the amide bond to generate the anhydride intermediate.

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AN 1997:225058 BIOSIS
DN PREV199799516774
TI Use of site-directed chemical modification to study an essential lysine in Escherichia coli leader peptidase.
AU Paetzel, Mark; Strynadka, Natalie C. J.; Tschantz, William R.; Casareno, Ruby; Bullinger, Patrick R.; Dalbey, Ross E. (1)
CS (1) Dep. Chem., Ohio State Univ., Columbus, OH 43210 USA
SO Journal of Biological Chemistry, (1997) Vol. 272, No. 15, pp. 9994-10003.
ISSN: 0021-9258.
DT Article

LA English

AB Escherichia coli leader peptidase, which catalyzes the cleavage of signal peptides from pre-proteins, is an essential, integral membrane serine peptidase that has its active site residing in the periplasmic space. It contains a conserved lysine residue that has been proposed to act as the general base, abstracting the proton from the side chain hydroxyl group of the nucleophilic serine 90. To help elucidate the role of the essential lysine 145 in the activity of E. coli leader peptidase, we have combined site-directed mutagenesis and chemical modification methods to introduce unnatural amino acid side chains at the 145-position. We show that partial activity can be restored to an inactive K145C leader peptidase mutant by reacting it with 2-bromoethylamine cntdot HBr to produce a lysine analog (y-thia-lysine) at the 145-position. Modification with the reagents 3-bromopropylamine cntdot HBr and 2-mercaptoethylamine also allowed for partial restoration of activity showing that there is some flexibility in the length requirements of this essential residue. Modification with (2-bromoethyl)trimethylammonium cntdot Br to form a positively charged, nontitratable side chain at the 145-position failed to restore activity to the inactive K145C leader peptidase mutant. This result, along with an inactive K145R mutant result, supports the claim that the lysine side chain at the 145-position is essential due to its ability to form a hydrogen bond(s) or to act as a general base rather than because of an ability to form a critical salt bridge. We find that leader peptidase processes the pre-protein substrate, pro-OmpA nuclease A, with maximum efficiency at pH 9.0, and apparent pK-a values for titratable groups at approximately 8.7 and 9.3 are revealed. We show that the lysine modifier maleic **anhydride** inhibits leader **peptidase** by reacting with lysine 145. The results of this study are consistent with the hypothesis that the lysine at the 145-position of leader peptidase functions as the active site general base. A model of the active site region of leader peptidase is presented based on the structure of the E. coli UmuD', and a mechanism for bacterial leader peptidase is proposed.

L2 ANSWER 31 OF 111 WPINDEX (C) 2003 THOMSON DERWENT

AN 1996-496428 [49] WPINDEX

DNC C1996-155107

TI Prepn. of valine derivs. HIV **protease** inhibitors - by converting mixed **anhydride** deriv. of N-((N-methyl-N-((2-isopropyl-4-thiazolyl)methyl)amino)carbonyl- valine to activated ester deriv..

DC B02 B03

IN COOPER, A J; MENZIA, J A; TIEN, J; TIEN, J J; TIEN, J H

PA (ABBO) ABBOTT LAB

CYC 22

PI US 5567823 A 19961022 (199649)* 7p
WO 9639398 A1 19961212 (199704) EN 22p
RW: AT BE CH DE DK ES FI FR GB GR IE IT LU MC NL PT SE
W: CA JP MX
EP 830353 A1 19980325 (199816) EN
R: AT BE CH DE DK ES FI FR GB GR IE IT LI LU NL PT SE
JP 11507029 W 19990622 (199935) 23p
MX 9709454 A1 19980201 (199954)
EP 830353 B1 20020424 (200228) EN
R: AT BE CH DE DK ES FI FR GB GR IE IT LI LU NL PT SE
DE 69620882 E 20020529 (200243)
ES 2176456 T3 20021201 (200305)

ADT US 5567823 A US 1995-469965 19950606; WO 9639398 A1 WO 1996-US6812 19960513; EP 830353 A1 EP 1996-915755 19960513, WO 1996-US6812 19960513; JP 11507029 W WO 1996-US6812 19960513, JP 1997-500554 19960513; MX 9709454 A1 MX 1997-9454 19971203; EP 830353 B1 EP 1996-915755 19960513, WO 1996-US6812 19960513; DE 69620882 E DE 1996-620882 19960513, EP 1996-915755 19960513, WO 1996-US6812 19960513; ES 2176456 T3 EP 1996-915755 19960513

FDT EP 830353 A1 Based on WO 9639398; JP 11507029 W Based on WO 9639398; EP 830353 B1 Based on WO 9639398; DE 69620882 E Based on EP 830353, Based on WO 9639398; ES 2176456 T3 Based on EP 830353

PRAI US 1995-469965 19950606

AB US 5567823 A UPAB: 19961205

Prepn. of(2S,3S,5S)-5-(N-(N-(N-methyl-N-((2-isopropyl-4-thiazolyl)methyl) amino)carbonyl-Dor-L-valinyl)amino)-2-(N-((5-thiazolyl) methoxycarbonyl(amino)-1,6-diphenyl-3-hydroxyhexane(I) or its acid addn. salt, comprises converting a mixed anhydride deriv. of N-((N-methyl-N-((2-isopropyl-4-thiazolyl)methyl)amino)carbonyl)(D or L-valine (II) to an activated ester deriv. and reacting this with (2S,3S,5S)-5-amino-2-)N-((S-thiazolyl)-methoxycarbonyl)amino)-1,6-diphenyl-3-hydroxyhexane (III).

USE - (I) are inhibitors of HIV-1 and HIV-2 protease.

Dwg.0/0

L2 ANSWER 35 OF 111 CAPLUS COPYRIGHT 2003 ACS

AN 1995:602399 CAPLUS

DN 123:47889

TI HIV protease inhibitors useful for the treatment of AIDS, and their preparation

IN Vacca, Joseph P.; Dorsey, Bruce D.; Guare, James P.; Holloway, M. Katharine; Hungate, Randall W.; Levin, Rhonda B.

PA Merck and Co., Inc., USA

SO U.S., 49 pp. Cont.-in-part of U.S. Ser. No. 40,729, abandoned. CODEN: USXXAM

DT Patent

LA English

FAN.CNT 5

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	US 5413999	A	19950509	US 1993-59038	19930507
	PL 171340	B1	19970430	PL 1992-303600	19921103
	RU 2131416	C1	19990610	RU 1994-27563	19921103
	RO 115726	B1	20000530	RO 1994-763	19921103
	CZ 287610	B6	20010117	CZ 1994-1110	19921103
	RU 2171254	C2	20010727	RU 1999-100203	19921103
	SK 281864	B6	20010806	SK 1994-523	19921103
	ZA 9208563	A	19930505	ZA 1992-8563	19921106
	BR 9406503	A	19960102	BR 1994-6503	19940324
	JP 08508496	T2	19960910	JP 1994-522189	19940324
	SK 279471	B6	19981104	SK 1995-1225	19940324
	RU 2139052	C1	19991010	RU 1995-122135	19940324
	RO 118000	B1	20021230	RO 1995-1690	19940324
	CA 2161334	AA	19941124	CA 1994-2161334	19940426
	WO 9426717	A1	19941124	WO 1994-US4621	19940426
	W: AU, BB, BG, BR, BY, CA, CN, CZ, FI, HU, JP, KR, KZ, LK, LV, MG, MN, MW, NO, NZ, PL, RO, RU, SD, SI, SK, TT, UA, UZ				
	RW: AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE, SN, TD, TG				
	AU 9466692	A1	19941212	AU 1994-66692	19940426
	AU 676563	B2	19970313		
	BR 9406576	A	19960130	BR 1994-6576	19940426
	EP 696277	A1	19960214	EP 1994-915427	19940426
	R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LI, LU, NL, PT, SE				
	HU 73135	A2	19960628	HU 1995-3170	19940426
	CN 1126469	A	19960710	CN 1994-192691	19940426
	JP 08509980	T2	19961022	JP 1994-525465	19940426
	ZA 9403104	A	19951106	ZA 1994-3104	19940505
	FI 9402112	A	19940506	FI 1994-2112	19940506
	NO 9401696	A	19940624	NO 1994-1696	19940506
	US 5527799	A	19960618	US 1995-407740	19950321

	FI 9504580	A	19950927	FI 1995-4580	19950927
	NO 9503876	A	19951130	NO 1995-3876	19950929
	FI 9505315	A	19951106	FI 1995-5315	19951106
	NO 9504427	A	19960108	NO 1995-4427	19951106
	US 5668132	A	19970916	US 1996-641720	19960502
	US 5717097	A	19980210	US 1996-759203	19961204
	CN 1176250	A	19980318	CN 1997-101853	19970201
	FI 9801591	A	19980710	FI 1998-1591	19980710
PRAI	US 1991-789508	B2	19911108		
	US 1992-883825	B2	19920515		
	US 1993-40729	B2	19930331		
	CS 1994-1110	A	19921103		
	WO 1992-US9444	W	19921103		
	US 1993-59038	A	19930507		
	WO 1994-US3209	W	19940324		
	WO 1994-US4621	W	19940426		
	US 1994-235576	B1	19940429		
	US 1995-407740	A3	19950321		
	US 1995-533142	B1	19950925		

OS MARPAT 123:47889

AB Compds. I [V = absent, C(O)Q, SO2Q (Q = absent, O, NR, (C1-4-substituted)heterocyclyl); R1 = (substituted) C1-4 alkyl, (substituted) aryl, (substituted) hetercyclyl, etc.; R3 = (substituted) benzyl; R12 = Q1, Q2] are claimed, as are compns. and methods for inhibiting HIV protease and treating AIDS. Prepn. of selected compds., e.g. N-[2(R)-hydroxy-1(S)-indanyl]-2(R)-phenylmethyl-4(S)-hydroxy-5-[1-(N'-(t-butyl)-4(S)-phenoxyprolineamide)yl]-pentaneamide, is described. IC50 values for selected compds. of the invention with respect to HIV protease inhibition are reported.

L2 ANSWER 39 OF 111 BIOTECHABS COPYRIGHT 2003 THOMSON DERWENT AND ISI

AN 1994-13792 BIOTECHABS

TI Protease chemical modification;
enzyme stabilization with an alkenyl ether and maleic anhydride copolymer, for use in protein hydrolysis

PA Nippon-Oil+Fats

PI JP 06205675 26 Jul 1994

AI JP 1991-65343 7 Mar 1991

PRAI JP 1991-65343 7 Mar 1991

DT Patent

LA Japanese

OS WPI: 1994-275517 [34]

AB A protease may be modified with a copolymer (I) which comprises an alkenyl ether, maleic anhydride and other monomers in a ratio of 5-60:20-90:0-30. In (I), Z is a residue with 2-8 OH groups, AO is a mixture of 1 or more 2-18C oxyalkylene groups (added in a block or at random), R1 is 2-5C alkenyl, R2 is 1-24C hydrocarbon or acyl, a, b and c are average addition molar numbers, each 0-600, m is 0-7, n is 0-6, m+n are 1-7, n/(1+m+n) is not more than 1/2, and a+bm+cn are 1-1,000. The modified protease, which is the reaction product of a copolymer (between an alkenyl ether with polyoxyalkylene groups and maleic **anhydride**) and **protease** possess no autolysis properties, and retain activity durably even in aq. solution. Use of the protease may be extended to the field of industrial protein hydrolysis. In an example, CH2=CHCH2O(C2H4O)33CH3 and maleic anhydride were polymerized to give 1,450 g copolymer, with a melting point of 45 deg and a saponification value of 68.5. The casein hydrolysis activity was 22 U660/mg, and residual activity was 72% after modification of subtilisin (EC-3.4.21.14). (9pp)

L2 ANSWER 55 OF 111 BIOSIS COPYRIGHT 2003 BIOLOGICAL ABSTRACTS INC.
DUPLICATE 9

AN 1992:365220 BIOSIS
DN BA94:47270
TI COUPLING OF DTPA TO PROTEINS A CRITICAL ANALYSIS OF THE CYCLIC DIANHYDRIDE METHOD IN THE CASE OF INSULIN MODIFICATION.
AU MAISANO F; GOZZINI L; DE HAEN C
CS BIOCHEMISTRY DEP., RESEARCH DEVELOPMENT DIVISION, BRACCO S.P.A., VIA E. FOLLI 50, 20134 MILAN, ITALY.
SO BIOCONJUGATE CHEM, (1992) 3 (3), 212-217.
CODEN: BCCHES. ISSN: 1043-1802.
FS BA; OLD
LA English
AB The reaction between the cyclic dianhydride of diethylenetriaminepentaacetic acid (DTPA), a bifunctional reagent, and proteins under various conditions was studied using porcine insulin as a model protein. The reaction was compared with that between citraconic anhydride, a monofunctional reagent, and insulin. Products were characterized chromatographically and electrophoretically before and after deesterification by hydroxylamine. A DTPA-conjugated product was further characterized by **proteolytic** fragmentation. The reaction with citraconic **anhydride** yielded the expected number of products exclusively acylated on amino groups. In contrast, the reaction with the cyclic dianhydride of DTPA under all conditions examined yielded a much higher number of products than expected. Among the products formed were O-acylated ones and products of intermolecular cross-linking. It is concluded that the use of the cyclic dianhydride of DTPA does not allow the reliable preparation of proteins or other macromolecules conjugated with a high number of DTPA molecules in which each molecule of DTPA is linked to one amino group of the macromolecule through a single amide bond.

L2 ANSWER 63 OF 111 BIOSIS COPYRIGHT 2003 BIOLOGICAL ABSTRACTS INC. DUPLICATE 13
AN 1988:502951 BIOSIS
DN BA86:123635
TI STUDYING CHEMICAL MODIFICATION OF PROTEOLYTIC ENZYMES WITH LOW-MOLECULAR WEIGHT AGENTS.
AU BEZ"YAZYCHNAYA T S; MOSKVICHEV B V
CS ALL-UNION RES. TECHNOL. INST. ANTIBIOT. ENZYMES MED. APPL., LENINGRAD, USSR.
SO PRIKL BIOKHIM MIKROBIOL, (1988) 24 (4), 481-483.
CODEN: PBMIAK. ISSN: 0555-1099.
FS BA; OLD
LA Russian
AB Low-molecular modification of **proteolytic** enzymes with aldehydes and **anhydrides** of carboxylic acids as well as with 2,4,6-trinitrobenzene sulphonic acid was studied. Specific activities of the enzymes were found to be dependent on the modification degree of their amino groups. The retaining of high activities in the region of low extents of enzyme modification enabled biocatalysts with activities similar to those of the native enzymes to be prepared.

L2 ANSWER 64 OF 111 CAPLUS COPYRIGHT 2003 ACS DUPLICATE 14
AN 1987:495276 CAPLUS
DN 107:95276
TI Preparation and chemical modification of microbial neutral proteases and their use as antitumor agents
IN Maeda, Hiroshi; Matsumura, Yasuhiro; Asami, Osamu; Tanaka, Hideyuki; Sasaki, Ikuharu
PA Amano Pharmaceutical Co., Ltd., Japan
SO Eur. Pat. Appl., 45 pp.
CODEN: EPXXDW
DT Patent

LA English
FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	EP 215662	A2	19870325	EP 1986-307088	19860915
	EP 215662	A3	19881019		
	R: AT, BE, CH, DE, FR, GB, IT, LI, LU, NL, SE				
	JP 62061926	A2	19870318	JP 1985-201607	19850913
	JP 06076339	B4	19940928		
	JP 63041426	A2	19880222	JP 1986-184126	19860807
	US 4844897	A	19890704	US 1986-906240	19860912
PRAI	JP 1985-201607		19850913		
	JP 1986-184126		19860807		

AB Microbial neutral proteases are shown to be effective anti-tumor agents, esp. after chem. modification, and they are formulated for use as medicaments. Proteases from S. marcescens (56K protease) and B. subtilis (AT protease) were prepd., chem. modified [e.g. with dextran, polyethylene glycol (PEG), succinate, methotrexate, cytosine arabinoside; crosslinked to form dimers, etc.], tested for cytotoxicity against various normal and tumor cells, and formulated. Tumor cells were selectively inhibited by the proteases, esp. in the presence of serum. Chem. modification had a significant effect on antitumor activity, e.g. measurement of the tumor vol. after various treatments showed (relative to addn. of the unmodified protease = 1) a vol. of 0.31 after addn. of AT-PEG and a vol. of 3.80 with no treatment.

L2 ANSWER 76 OF 111 CAPLUS COPYRIGHT 2003 ACS DUPLICATE 18
AN 1982:158255 CAPLUS
DN 96:158255
TI Stabilization of microbial proteases against autolysis using acylation with dicarboxylic acid anhydrides
AU Maneepun, Saipin; Klibanov, Alexander M.
CS Dep. Nutr. Food Sci., Massachusetts Inst. Technol., Cambridge, MA, 02139, USA
SO Biotechnology and Bioengineering (1982), 24(2), 483-6
CODEN: BIBIAU; ISSN: 0006-3592
DT Journal
LA English
AB Immobilization of Streptomyces caespitosus or Bacillus thermoproteolyticus proteinases on CNBr-activated Sepharose markedly decreased the rate of inactivation obsd. upon incubation of the free enzymes at 45.degree.. Modification of the B. thermoproteolyticus **proteinase** with succinic or malic **anhydrides** prevented autolysis and decreased thermoinactivation. Acetylation also prevented autolysis.

L2 ANSWER 82 OF 111 BIOSIS COPYRIGHT 2003 BIOLOGICAL ABSTRACTS INC.
AN 1979:160943 BIOSIS
DN BA67:40943
TI REACTION OF A MIXED ANHYDRIDE WITH AQUEOUS HYDROXYLAMINE A MODEL FOR THE TRAPPING BY ADDED NUCLEOPHILES OF **ANHYDRIDE** INTERMEDIATES IN CARBOXY **PEPTIDASE** A ACTION.
AU SUGIMOTO T; KAISER E T
CS DEP. CHEM., UNIV. CHIC., CHICAGO, ILL. 60637, USA.
SO J ORG CHEM, (1978) 43 (17), 3311-3313.
CODEN: JOCEAH. ISSN: 0022-3263.
FS BA; OLD
LA English
AB As a model for experiments on the trapping by nucleophiles of acyl-enzyme intermediates formed in the action of carboxpeptidase A, the reaction of trans-p-chlorocinnamic propionic anhydride with aqueous hydroxylamine was examined. Both above and below the pKa of hydroxylamine, propionohydroxamic acid was formed in very high yields. The other dominant

product was trans-p-chlorocinnamic acid. The pH-rate constant profile for the attack of hydroxylamine on the mixed anhydride was sigmoidal, with an apparent pKa value of 6.07 .+-. 0.11 and a limiting 2nd-order rate constant of 2340 M-1 s-1 calculated in alkaline solution. Within the limits of measurement, catalysis of anhydride breakdown occurred only with the unprotonated form of hydroxylamine. The results suggest that if the acyl-enzyme intermediate observed in kinetic measurements on the reaction of carboxypeptidase A with O-(trans-p-chlorocinnamoyl)-L-.beta.-phenyllactate is an anhydride species, nucleophilic trapping with hydroxylamine in the absence of interaction of the active site metal ion with the anhydride may be accomplished in reasonable yields.

L2 ANSWER 91 OF 111 CAPLUS COPYRIGHT 2003 ACS DUPLICATE 22
AN 1971:415311 CAPLUS
DN 75:15311
TI Fixation of **proteolytic** enzymes on poly(methacrylic **anhydride**)
AU Conte, Apollonio; Lehmann, Klaus
CS Pharm. Lab., Roehm G.m.b.H., Darmstadt, Fed. Rep. Ger.
SO Hoppe-Seyler's Zeitschrift fuer Physiologische Chemie (1971), 352(4), 533-41
CODEN: HSZPAZ; ISSN: 0018-4888
DT Journal
LA German
AB Enzymically active ensyme resins were prepd. by fixing the proteolytic enzymes, trypsin, chymotrypsin, and papain to cross-linked poly(methacrylic anhydride). These prepns. retained 3-20% of the enzymic activity toward casein and .ltoreq.40% toward low mol. wt. substrates, compared with the free enzyme. The binding of the enzymes to the carrier resulted in a considerable stabilization of enzyme activity. These resins could be used many times without appreciable loss of activity.

L2 ANSWER 96 OF 111 CAPLUS COPYRIGHT 2003 ACS DUPLICATE 23
AN 1969:487907 CAPLUS
DN 71:87907
TI Identification of lysine and arginine residues as inhibitory centers of protease inhibitors with the aid of maleic anhydride and 2,3-butandione
AU Fritz, Hans; Fink, Edwin; Gebhardt, Maria; Hochstrasser, Karl; Werle, Eugen
CS Univ. Muenchen, Munich, Fed. Rep. Ger.
SO Hoppe-Seyler's Zeitschrift fuer Physiologische Chemie (1969), 350(8), 933-44
CODEN: HSZPAZ; ISSN: 0018-4888
DT Journal
LA German
AB The following inhibitors were treated with maleic anhydride, whereupon they lost their inhibitory activity towards the given enzymes: inhibitor from swine, dog, and cat pancreas (trypsin); from guinea pig seminal vesicles, hirudin, and lima beans (trypsin and plasmin); from bovine and sheep lung (trypsin, plasmin, kallikrein, and chymotrypsin). The antichymotryptic activity of the inhibitor from lima beans was not affected by acylation of the amino groups. The inhibitors regained their inhibitory activity after deacylation in acidic soln. The polymaleoyl derivs. of the inhibitors from haricot beans and guinea pig seminal vesicles still possessed .apprx.1/3 of the antitryptic activity of the native inhibitors. The loss of inhibition towards trypsin or plasmin and kallikrein after reaction with maleic anhydride is due to the acylation of the amino group of a lysine residue, which is in the reactive center of the inhibitor. The following inhibitors contain an arginine residue in the reactive center: inhibitor from sheep pancreas, from submandibular gland of the dog, from soybean, from hen egg white, wheat shoots, rye shoots, potatoes, ground nuts, and the inter-.alpha.-trypsin inhibitor

from human serum. The polymaleoyl derivs. of these inhibitors, which possess the same antitryptic activity as the native inhibitors, are inactivated irreversibly and relatively quickly by reaction with a 2,3-butanedione reagent. This reagent modifies specifically the guanidino groups of the arginine residues after acylation of the amino groups of the inhibitors. The antiplasmin activity of the inhibitors from the submandibular gland of the dog, soybean, and ground nuts is not decreased after the acylation of the amino groups, but when the polymaleoyl derivs. of these inhibitors are treated with 2,3-butanedione reagent, the decrease of their antiplasmin activities parallels that of their antitrypsin activities.

L2 ANSWER 104 OF 111 SYNTHLINE COPYRIGHT 2003 PROUS SCIENCE
AN 2000:3099 SYNTHLINE
TI Symmetrical **anhydride**-type serine **protease** inhibitors: Structure-activity relationship studies of human chymase inhibitors
AU Katada, J.; Hayashi, Y.; Iijima, K.
SO Bioorg Med Chem Lett (1999), 9(3), 413

=>
<------User Break------>

L Number	Hits	Search Text	DB	Time stamp
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NEWS 8	Sep 16	Experimental properties added to the REGISTRY file
NEWS 9	Sep 16	CA Section Thesaurus available in CAPLUS and CA
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NEWS 15	Dec 04	CSA files on STN
NEWS 16	Dec 17	PCTFULL now covers WP/PCT Applications from 1978 to date
NEWS 17	Dec 17	TOXCENTER enhanced with additional content
NEWS 18	Dec 17	Adis Clinical Trials Insight now available on STN
NEWS 19	Jan 29	Simultaneous left and right truncation added to COMPENDEX, ENERGY, INSPEC
NEWS 20	Feb 13	CANCERLIT is no longer being updated
NEWS 21	Feb 24	METADEX enhancements
NEWS 22	Feb 24	PCTGEN now available on STN
NEWS 23	Feb 24	TEMA now available on STN
NEWS 24	Feb 26	NTIS now allows simultaneous left and right truncation
NEWS 25	Feb 26	PCTFULL now contains images
NEWS 26	Mar 04	SDI PACKAGE for monthly delivery of multifile SDI results
NEWS 27	Mar 20	EVENTLINE will be removed from STN
NEWS 28	Mar 24	PATDPAFULL now available on STN
NEWS 29	Mar 24	Additional information for trade-named substances without structures available in REGISTRY
NEWS 30	Apr 11	Display formats in DGENE enhanced
NEWS 31	Apr 14	MEDLINE Reload
NEWS 32	Apr 17	Polymer searching in REGISTRY enhanced
NEWS 33	Jun 13	Indexing from 1947 to 1956 added to records in CA/CAPLUS
NEWS 34	Apr 21	New current-awareness alert (SDI) frequency in WPIDS/WPINDEX/WPIX
NEWS 35	Apr 28	RDISCLOSURE now available on STN
NEWS 36	May 05	Pharmacokinetic information and systematic chemical names added to PHAR
NEWS 37	May 15	MEDLINE file segment of TOXCENTER reloaded
NEWS 38	May 15	Supporter information for ENCOMPPAT and ENCOMPLIT updated
NEWS 39	May 16	CHEMREACT will be removed from STN
NEWS 40	May 19	Simultaneous left and right truncation added to WSCA
NEWS 41	May 19	RAPRA enhanced with new search field, simultaneous left and right truncation
NEWS 42	Jun 06	Simultaneous left and right truncation added to CBNB

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  NEWS 43  Jun 06  PASCAL enhanced with additional data
  NEWS 44  Jun 20  2003 edition of the FSTA Thesaurus is now available
  NEWS 45  Jun 25  HSDB has been reloaded

  NEWS EXPRESS  April 4 CURRENT WINDOWS VERSION IS V6.01a, CURRENT
                MACINTOSH VERSION IS V6.0b(ENG) AND V6.0Jb(JP),
                AND CURRENT DISCOVER FILE IS DATED 01 APRIL 2003
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 NEWS 44  Jun 20  2003 edition of the FSTA Thesaurus is now available
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=> s (blood (2A) factor) (3A) (anhydridize or anhydride or anhydridized or
anhyddridization)
  12 FILES SEARCHED...
  22 FILES SEARCHED...
  30 FILES SEARCHED...
  43 FILES SEARCHED...
  55 FILES SEARCHED...
  67 FILES SEARCHED...
  87 FILES SEARCHED...
 103 FILES SEARCHED...
L1          10 (BLOOD (2A) FACTOR) (3A) (ANHYDRIDIZE OR ANHYDRIDE OR ANHYDRIDIZ
               ED OR ANHYDDRIDIZATION)

=> s (blood (2A) factor) (3A) (anhydridization)
  13 FILES SEARCHED...
  31 FILES SEARCHED...
  48 FILES SEARCHED...
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  96 FILES SEARCHED...
L2           0 (BLOOD (2A) FACTOR) (3A) (ANHYDRIDIZATION)

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HSDB, MSDS-CCOHS, MSDS-OHS, RTECS, CONF, IMSDRUGCONF, DIOGENES, INVESTEXT,
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L3   ANSWER 1 OF 10  CAPLUS  COPYRIGHT 2003 ACS
AN   2003:5928  CAPLUS
DN   138:73271
TI   Preparation of N,N'-bis(heterocyclic acyl)cycloalkanediamine and
     heterocyclediamine derivatives as inhibitors of activated blood
     coagulation factor X (factor Xa)
IN   Ohta, Toshiharu; Komoriya, Satoshi; Yoshino, Toshiharu; Uoto, Kouichi;
     Nakamoto, Yumi; Naito, Hiroyuki; Mochizuki, Akiyoshi; Nagata, Tsutomu;
     Kanno, Hideyuki; Haginoya, Noriyasu; Yoshikawa, Kenji; Nagamochi,
     Masatoshi; Kobayashi, Syozo; Ono, Makoto
PA   Daiichi Pharmaceutical Co., Ltd., Japan
SO   PCT Int. Appl., 788 pp.
     CODEN: PIXXD2
DT   Patent
LA   Japanese
FAN.CNT 3
     PATENT NO.        KIND  DATE           APPLICATION NO.  DATE
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PI   WO 2003000657     A1    20030103       WO 2002-JP2683   20020320
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             CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH,
             GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,
             LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH,
             PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ,
             UA, UG, US, UZ, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU,
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WO 2003000680 A1 20030103 WO 2002-JP6141 20020620
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
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WO 2003016302 A1 20030227 WO 2002-JP8119 20020808
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
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PRAI JP 2001-187105 A 20010620
JP 2001-243046 A 20010809
JP 2001-311808 A 20011009
JP 2001-398708 A 20011228
WO 2002-JP2683 W 20020320
WO 2002-JP6141 A 20020620
OS MARPAT 138:73271
RE.CNT 54 THERE ARE 54 CITED REFERENCES AVAILABLE FOR THIS RECORD
ALL CITATIONS AVAILABLE IN THE RE FORMAT

L3 ANSWER 2 OF 10 USPATFULL
AN 2003:106233 USPATFULL
TI Compositions and methods for the therapy and diagnosis of pancreatic cancer
IN Benson, Darin R., Seattle, WA, UNITED STATES
Kalos, Michael D., Seattle, WA, UNITED STATES
Lodes, Michael J., Seattle, WA, UNITED STATES
Persing, David H., Redmond, WA, UNITED STATES
Hepler, William T., Seattle, WA, UNITED STATES
Jiang, Yuqiu, Kent, WA, UNITED STATES
PA Corixa Corporation, Seattle, WA, UNITED STATES, 98104 (U.S. corporation)
PI US 2003073144 A1 20030417
AI US 2002-60036 A1 20020130 (10)
PRAI US 2001-333626P 20011127 (60)
US 2001-305484P 20010712 (60)
US 2001-265305P 20010130 (60)
US 2001-267568P 20010209 (60)
US 2001-313999P 20010820 (60)
US 2001-291631P 20010516 (60)
US 2001-287112P 20010428 (60)
US 2001-278651P 20010321 (60)
US 2001-265682P 20010131 (60)
DT Utility
FS APPLICATION
LREP SEED INTELLECTUAL PROPERTY LAW GROUP PLLC, 701 FIFTH AVE, SUITE 6300, SEATTLE, WA, 98104-7092
CLMN Number of Claims: 17

ECL Exemplary Claim: 1
DRWN No Drawings
LN.CNT 14253
CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L3 ANSWER 3 OF 10 USPATFULL
AN 2002:272801 USPATFULL
TI Compositions and methods for the therapy and diagnosis of colon cancer
IN Stolk, John A., Bothell, WA, UNITED STATES
Xu, Jiangchun, Bellevue, WA, UNITED STATES
Chenault, Ruth A., Seattle, WA, UNITED STATES
Meagher, Madeleine Joy, Seattle, WA, UNITED STATES
PA Corixa Corporation, Seattle, WA, UNITED STATES, 98104 (U.S. corporation)
PI US 2002150922 A1 20021017
AI US 2001-998598 A1 20011116 (9)
PRAI US 2001-304037P 20010710 (60)
US 2001-279670P 20010328 (60)
US 2001-267011P 20010206 (60)
US 2000-252222P 20001120 (60)
DT Utility
FS APPLICATION
LREP SEED INTELLECTUAL PROPERTY LAW GROUP PLLC, 701 FIFTH AVE, SUITE 6300, SEATTLE, WA, 98104-7092
CLMN Number of Claims: 17
ECL Exemplary Claim: 1
DRWN No Drawings
LN.CNT 9233
CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L3 ANSWER 4 OF 10 USPATFULL
AN 2002:243051 USPATFULL
TI Compositions and methods for the therapy and diagnosis of ovarian cancer
IN Algate, Paul A., Issaquah, WA, UNITED STATES
Jones, Robert, Seattle, WA, UNITED STATES
Harlocker, Susan L., Seattle, WA, UNITED STATES
PA Corixa Corporation, Seattle, WA, UNITED STATES, 98104 (U.S. corporation)
PI US 2002132237 A1 20020919
AI US 2001-867701 A1 20010529 (9)
PRAI US 2000-207484P 20000526 (60)
DT Utility
FS APPLICATION
LREP SEED INTELLECTUAL PROPERTY LAW GROUP PLLC, 701 FIFTH AVE, SUITE 6300, SEATTLE, WA, 98104-7092
CLMN Number of Claims: 11
ECL Exemplary Claim: 1
DRWN No Drawings
LN.CNT 25718
CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L3 ANSWER 5 OF 10 CAPLUS COPYRIGHT 2003 ACS
AN 2000:900684 CAPLUS
DN 134:46756
TI Substance binding to the substrate of activated blood coagulation factor in competition with this factor to thereby regulate the reaction between the activated blood coagulation factor and the substrate, a process for producing the substance and blood coagulation factor-adsorbent with the use of the substance
IN Hosokawa, Kazuya
PA Fujimori Kogyo Co., Ltd., Japan; Chisso Corporation
SO PCT Int. Appl., 24 pp.
CODEN: PIXXD2
DT Patent
LA Japanese
FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO 2000077048	A1	20001221	WO 2000-JP3863	20000614
	W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
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	EP 1188770	A1	20020320	EP 2000-937225	20000614
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PRAI	JP 1999-167453	A	19990614		
	JP 2000-62629	A	20000307		
	WO 2000-JP3863	W	20000614		

RE.CNT 7 THERE ARE 7 CITED REFERENCES AVAILABLE FOR THIS RECORD
ALL CITATIONS AVAILABLE IN THE RE FORMAT

L3 ANSWER 6 OF 10 CAPLUS COPYRIGHT 2003 ACS
AN 1999:529128 CAPLUS
DN 131:184864
TI Preparation of amidinophenylcarbamoylbiphenyl derivatives and heterocyclic analogs thereof as inhibitors of blood coagulation factor VIIa
IN Senokuchi, Kazuhiko; Ogawa, Koji
PA Ono Pharmaceutical Co., Ltd., Japan
SO PCT Int. Appl., 665 pp.
CODEN: PIXXD2
DT Patent
LA Japanese
FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO 9941231	A1	19990819	WO 1999-JP622	19990212
	W: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GE, GH, GM, HR, HU, ID, IL, IS, JP, KE, KG, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
	RW: GH, GM, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG				
	AU 9923006	A1	19990830	AU 1999-23006	19990212
	EP 1078917	A1	20010228	EP 1999-902896	19990212
	R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, FI				
	ZA 9901273	A	19990825	ZA 1999-1273	19990217
	US 6358960	B1	20020319	US 2000-601998	20000811
PRAI	JP 1998-76815	A	19980217		
	WO 1999-JP622	W	19990212		

OS MARPAT 131:184864
RE.CNT 5 THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS RECORD
ALL CITATIONS AVAILABLE IN THE RE FORMAT

L3 ANSWER 7 OF 10 CAPLUS COPYRIGHT 2003 ACS
AN 1989:91686 CAPLUS
DN 110:91686
TI Antigenic analogs of platelet-activating factor (PAF), production of the analogs and antibodies to them, and PAF immunoassays
IN Baldo, Brian Angelo; Redmond, John William
PA University of Sydney, Australia; Macquarie University; Royal North Shore Hospital

SO PCT Int. Appl., 46 pp.
CODEN: PIXXD2
DT Patent
LA English
FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO 8705904	A1	19871008	WO 1987-AU84	19870324
	W: AU, JP, KR, US				
	RW: DE, FR, GB, IT				
	AU 8772097	A1	19871020	AU 1987-72097	19870324
	AU 607698	B2	19910314		
	EP 299965	A1	19890125	EP 1987-902318	19870324
	R: DE, FR, GB, IT				
	JP 01502584	T2	19890907	JP 1987-502157	19870324
	IL 82057	A1	19941111	IL 1987-82057	19870331
	US 5061626	A	19911029	US 1987-156923	19871124
PRAI	AU 1986-5175		19860324		
	WO 1987-AU84		19870324		

L3 ANSWER 8 OF 10 CAPLUS COPYRIGHT 2003 ACS
AN 1978:419780 CAPLUS
DN 89:19780
TI Separating a factor IX preparation from plasma using ethylene-maleic anhydride polymers
IN Delente, Jacques J.; Schoenfeld, Richard A.
PA Monsanto Co., USA
SO U.S., 4 pp.
CODEN: USXXAM
DT Patent
LA English
FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	US 4081432	A	19780328	US 1977-818920	19770725
	ES 471857	A1	19790201	ES 1978-471857	19780719
	EP 651	A1	19790207	EP 1978-300177	19780721
	EP 651	B1	19820127		
	R: BE, CH, DE, FR, GB, NL, SE				
	JP 54026322	A2	19790227	JP 1978-89306	19780721
	JP 61054008	B4	19861120		
	AU 7838243	A1	19800124	AU 1978-38243	19780721
	AU 517885	B2	19810903		
	AT 7805317	A	19800415	AT 1978-5317	19780721
	AT 359645	B	19801125		
	RO 75338	P	19801130	RO 1978-94744	19780721
	SU 841572	A3	19810623	SU 1978-2639950	19780721
	IL 55194	A1	19810731	IL 1978-55194	19780721
	HU 23516	O	19820928	HU 1978-MO1022	19780721
	HU 180882	B	19830530		
	CA 1107649	A1	19810825	CA 1978-307999	19780724
PRAI	US 1977-818920		19770725		

L3 ANSWER 9 OF 10 CAPLUS COPYRIGHT 2003 ACS
AN 1977:186766 CAPLUS
DN 86:186766
TI Products of the citraconylation of bull prothrombin and their activation by factor X
AU Memon, M. S.; Baskova, I. P.
CS Lab. Fiziol. Biokhim. Svertyvaniya Krovi, Mosk. Gos. Univ. im. Lomonosova, Moscow, USSR
SO Biokhimiya (Moscow) (1977), 42(3), 505-12
CODEN: BIOHAO; ISSN: 0320-9725
DT Journal

LA Russian

L3 ANSWER 10 OF 10 DDFB COPYRIGHT 2003 THOMSON DERWENT
AN 1972-13036 DDFB P X
TI FACTOR VIII /AHF/ ACTIVITY OF SMALL SIZE PRODUCED BY SUCCINYLATING PLASMA.
AU BARROW E M; GRAHAM J B
LO CHAPEL HILL,N.C.
SO AM.J.PHYSIOL. (222, NO.1, 134-41, 1972)
DT Journal

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FULL ESTIMATED COST	10.50	766.45

DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)	SINCE FILE ENTRY	TOTAL SESSION
CA SUBSCRIBER PRICE	0.00	-0.65

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